



ARCD:307USD1 09/768,877

## **APPENDIX A:**

## PENDING CLAIMS AS OF SECOND RESTRICTION REQUIREMENT

- 18. A method of screening for a modulator of calpain function comprising:
  - a) obtaining a calpain polypeptide;
  - b) determining a standard activity profile of the calpain polypeptide;
  - c) contacting the calpain polypeptide with a putative modulator; and
  - d) assaying for a change in the standard activity profile.

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- 19. The method of claim 18, wherein the calpain polypeptide is a calpain 10 polypeptide.
- 20. The method of claim 18, wherein obtaining the calpain polypeptide comprises expressing the polypeptide in a host cell.
- 21. The method of claim 20, wherein the calpain polypeptide is isolated away from the host cell prior to contacting the calpain polypeptide with the putative modulator.
- 49. The method of claim 19, wherein the standard activity profile of the calpain 10 polypeptide is determined by measuring the binding of the calpain 10 polypeptide to a synthetic substrate.
- 50. The method of claim 49, wherein the synthetic substrate is Suc-Leu-Tyr-AMC.
- 51. A method of screening for a modulator of calpain function comprising:
  - a) obtaining an calpain polypeptide;
  - b) contacting the calpain polypeptide with a putative modulator; and
  - c) assaying for modulation of calpain function by the putative modulator.
- 52. The method of claim 51, wherein the calpain polypeptide is a calpain 10 polypeptide.

- The method of claim 52, wherein the calpain 10 polypeptide has a sequence comprising SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:6, SEQ ID NO:8, SEQ ID NO:10, SEQ ID NO:12, SEQ ID NO:14, SEQ ID NO:16, or SEQ ID NO:18.
- 54. The method of claim 51, further comprising determining a standard activity profile of the calpain polypeptide.
- 55. The method of claim 54, wherein the standard activity profile of the calpain 10 polypeptide is determined by measuring the binding of the calpain 10 polypeptide to a synthetic substrate.
- 56. The method of claim 55, wherein the synthetic substrate is Suc-Leu-Tyr-AMC.
- 57. The method of claim 55, wherein assaying for modulation of calpain function comprises assaying for a change in the standard activity profile.
- 58. The method of claim 51, wherein obtaining the calpain polypeptide comprises expressing the polypeptide in a host cell.
- 59. The method of claim 58, wherein the calpain polypeptide is isolated away from the host cell prior to contacting the calpain polypeptide with the putative modulator.
- 60. The method of claim 51, wherein obtaining the calpain polypeptide comprises obtaining a cell containing the polypeptide.
- 61. The method of claim 60, wherein the cell is a pancreatic cell, a muscle cell, an adipose cell, or a liver cell.
- 62. The method of claim 61, wherein the cell is a pancreatic cell.

- 63. The method of claim 62, wherein the pancreatic cell is comprised in an isolated pancreatic islet.
- 64. The method of claim 62, wherein the cell is a  $\beta$ -cell.